

Application Serial No.: 10/088,744  
Attorney Docket No.: 01975-0034-00000

**IN THE CLAIMS:**

Please amend claims 1, 2, 5-10, 12-25, 28-30, 33-36, 39, and 40, cancel claims 3 and 4, and add new claims 41-69, as follows:

1. (CURRENTLY AMENDED) An isolated polynucleotide comprising a nucleotide sequence ~~selected from the group consisting of~~ chosen from:
  - a) a nucleotide sequence of SEQ ID NO: 1, SEQ ID NO: 3, SEQ ID NO: 5 or SEQ ID NO: 7;
  - a)b) a nucleotide sequence encoding the IGS4 polypeptide according to SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6 or SEQ ID NO: 8.;
  - b)c) a nucleotide sequence encoding the polypeptide encoded by the DNA insert contained in the deposit no. CBS102221 or the deposit no. CBS102222 at the Centraalbureau voor Schimmelcultures at Baarn the Netherlands, ~~in particular a nucleotide sequence corresponding to the SEQ ID NO: 1, SEQ ID NO: 3, SEQ ID NO: 5 or SEQ ID NO: 7.~~
  - e)d) a nucleotide sequence having at least 80 % (~~preferably at least 90%~~) sequence identity over its entire length to the nucleotide sequence of (a), ~~or (b), or (c); or~~
  - d)e) a nucleotide sequence which is complimentary to the nucleotide sequence of (a) or (b) or (c) or (d).
2. (CURRENTLY AMENDED) The polynucleotide of claim 1 wherein said polynucleotide comprises:

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a) the nucleotide sequence contained in SEQ ID NO: 1 encoding the IGS4 polypeptide of SEQ ID NO: 2; or

b) the nucleotide sequence contained in SEQ ID NO: 3 encoding the IGS4 polypeptide of SEQ ID NO: 4; or

c) the nucleotide sequence contained in SEQ ID NO: 5 encoding the IGS4 polypeptide of SEQ ID NO: 6; or

d) the nucleotide sequence contained in SEQ ID NO: 7 encoding the IGS4 polypeptide of SEQ ID NO: 8.

3. (CANCELED)

4. (CANCELED)

5. (CURRENTLY AMENDED) The polynucleotide of ~~claim 1-4~~ claim 1 which is DNA or RNA.

6. (CURRENTLY AMENDED) An isolated nucleotide sequence encoding an IGS4 neuromedin receptor protein, ~~preferably a mammalian neuromedin receptor protein~~, said protein exhibiting high affinity binding for neuromedin U, ~~preferably for neuromedin U-8, for neuromedin U-23 and/or for neuromedin U-25.~~

7. (CURRENTLY AMENDED) An The isolated nucleotide sequence of claim 6 encoding an IGS4 neuromedin receptor protein, said protein exhibiting expression in at least one of brain, skeletal muscle, cerebellum, testis, corpus callosum, spinal cord, substantia nigra, medulla, thalamus, caudate nucleus,

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pons, nucleus accumbens, fetal brain, stomach, heart, thyroid gland, lung,  
thymus, prostate, and and/or in trachea.

8. (CURRENTLY AMENDED) An isolated nucleotide sequence encoding an IGS4 neuromedin receptor protein, ~~preferably a mammalian neuromedin receptor protein~~, said protein exhibiting high affinity binding for neuromedin U, ~~preferably for neuromedin U-8, for neuromedin U-23 and/or for neuromedin U-25~~, said protein exhibiting expression in at least one of brain, skeletal muscle, cerebellum, testis, corpus callosum, spinal cord, substantia nigra, medulla, thalamus, caudate nucleus, pons, nucleus accumbens, fetal brain, stomach, heart, thyroid gland, lung, thymus, prostate, and and/or in trachea, and said nucleotide sequence being selected from the group of nucleotide sequences as defined in ~~the claims 4 to claim 5~~ claim 1.
9. (CURRENTLY AMENDED) An expression system comprising a A DNA or RNA molecule ~~comprising an expression system~~, wherein said expression system is ~~capable of producing~~ produces an IGS4 polypeptide comprising an amino acid sequence, which has at least 80% identity with the polypeptide of SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 6 or SEQ ID NO: 8 or with the polypeptide encoded by the DNA insert contained in the deposit no. CBS1 02221 or the deposit no. CBS1 02222 at the Centraalbureau voor Schimmelcultures at Baarn the Netherlands, when said expression system is present in a compatible host cell.
10. (CURRENTLY AMENDED) An expression system comprising an An isolated DNA or RNA molecule ~~comprising an expression system~~, wherein said

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expression system ~~is capable of producing~~ produces an IGS4 polypeptide comprising an amino acid sequence which is a neuromedin receptor protein, ~~preferably a mammalian neuromedin receptor protein~~, said protein exhibiting high affinity binding for neuromedin U, ~~preferably for neuromedin U-8, for neuromedin-U-23 and/or for neuromedin-U-25~~, and exhibiting expression in at least one of brain, skeletal muscle, cerebellum, testis, corpus callosum, spinal cord, substantia nigra, medulla, thalamus, caudate nucleus, pons, nucleus accumbens, fetal brain, stomach, heart, thyroid gland, lung, thymus, prostate, and ~~and/or in~~ trachea.

11. (CURRENTLY AMENDED) A host cell comprising the expression system of claim 9 or 10.
12. (CURRENTLY AMENDED) ~~A~~ The host cell according to claim 11 ~~which~~ wherein the host cell is a yeast cell.
13. (CURRENTLY AMENDED) A host cell according to claim 11 ~~which~~ wherein the host cell is an animal cell.
14. (CURRENTLY AMENDED) An IGS4 receptor membrane preparation ~~derived~~ prepared from a cell according to claim 11-13.
15. (CURRENTLY AMENDED) A process for producing an IGS4 polypeptide comprising culturing a host of claim 11-13 under conditions sufficient for the production of said polypeptide and recovering the polypeptide from the culture.

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16. (ORIGINAL) A process for producing a cell which produces an IGS4 polypeptide comprising transforming or transfecting a host cell with the expression system of claim 9 such that the host cell, under appropriate culture conditions, produces an IGS4 polypeptide.
17. (CURRENTLY AMENDED) An IGS4 polypeptide comprising an amino acid sequence ~~which is at least 80% identical to the amino acid sequence~~ of SEQ ID NO: 2, SEQ NO: 4, SEQ NO: 6 or SEQ NO: 8 or ~~to~~ the polypeptide encoded by the DNA insert contained in the deposit no. CBS102221 or the deposit no. CBS102222 at the Centraalbureau voor Schimmelcultures at Baarn the Netherlands over its entire length.
18. (CURRENTLY AMENDED) The polypeptide of claim 17 ~~which~~ wherein the polypeptide is at least 80% identical to ~~comprises~~ the amino acid sequence of SEQ ID NO: 2, SEQ ID NO: 4, SEQ NO: 6 or SEQ NO: 8 or the amino acid sequence encoded by the DNA insert contained in the deposit no. CBS102221 or the deposit no. CBS102222 at the Centraalbureau voor Schimmelcultures at Baarn the Netherlands.
19. (CURRENTLY AMENDED) An isolated IGS4 polypeptide comprising an amino acid sequence of a neuromedin receptor protein, ~~preferably of a mammalian neuromedin receptor protein~~, said protein exhibiting high affinity binding for neuromedin U, ~~preferably for neuromedin U-8, for neuromedin U-23 and/or for neuromedin U-25.~~

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20. (CURRENTLY AMENDED) An The isolated IGS4 polypeptide of claim 19 comprising an amino acid sequence of a neuromedin receptor protein, said protein exhibiting expression in at least one of brain, skeletal muscle, cerebellum, testis, corpus callosum, spinal cord, substantia nigra, medulla, thalamus, caudate nucleus, pons, nucleus accumbens, fetal brain, stomach, heart, thyroid gland, lung, thymus, prostate, and ~~and/or~~ in trachea.
21. (CURRENTLY AMENDED) An isolated IGS4 polypeptide comprising an amino acid sequence of a neuromedin receptor protein, ~~preferably a mammalian neuromedin receptor protein~~, said protein exhibiting high affinity binding for neuromedin U, ~~preferably for neuromedin U-8, for neuromedin U-23 and/or for neuromedin U-26~~, said protein exhibiting expression in at least one of brain, skeletal muscle, cerebellum, testis, corpus callosum, spinal cord, substantia nigra, medulla, thalamus, caudate nucleus, pons, nucleus accumbens, fetal brain, stomach, heart, thyroid gland, lung, thymus, prostate, and ~~and/or~~ in trachea, and said amino acid sequence being selected from the group of amino acid sequence sequences as defined in the ~~claims 17-18~~ claim 17.
22. (CURRENTLY AMENDED) An antibody immunospecific for the IGS4 polypeptide of claim 17-24.
23. (CURRENTLY AMENDED) A method for the treatment of a subject in need of enhanced activity or expression of IGS4 polypeptide of claim 17-24 comprising at least one of:

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(a) administering to the subject a therapeutically effective amount of an agonist to said receptor; ~~and/or~~

(b) providing to the subject an isolated polynucleotide comprising a nucleotide sequence that has at least 80% identity to a nucleotide sequence encoding the IGS4 polypeptide of SEQ ID NO: 2, SEQ ID NO: 4, SEQ NO: 6 or SEQ NO: 8 or the polypeptide encoded by the DNA insert contained in the deposit no. CBS102221 or the deposit no. CBS102222 at the Centraalbureau voor Schimmelcultures at Baarn the Netherlands over its entire length; or a nucleotide sequence complementary to one of said nucleotide sequences in a form so as to effect production of said receptor activity in vivo; and,

(c) providing to the subject an isolated polynucleotide comprising a nucleotide sequence that encodes an IGS4 neuromedin receptor protein, ~~preferably a mammalian IGS4 neuromedin receptor protein~~, said protein exhibiting high affinity binding for neuromedin U, ~~preferably for neuromedin U-8, for neuromedin U-23 and/or for neuromedin U-25.~~

24. (CURRENTLY AMENDED) A method for the treatment of a subject having need to inhibit activity or expression of IGS4 polypeptide of claim 17-~~24~~ comprising at least one of:

(a) administering to the subject a therapeutically effective amount of an antagonist to said receptor; ~~and/or~~

(b) administering to the subject a nucleic acid molecule that inhibits the expression of the nucleotide sequence encoding said receptor; ~~and/or~~ and,

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(c) administering to the subject a therapeutically effective amount of a polypeptide that competes with said receptor for its ligand.

25. (CURRENTLY AMENDED) A process for diagnosing a disease or a susceptibility to a disease in a subject related to expression or activity of the IGS4 polypeptide of claim 17-24 in a subject comprising at least one of:

(a) determining the presence or absence of a mutation in the nucleotide sequence encoding said IGS4 polypeptide in the genome of said subject; ~~and/or~~ and,

(b) analyzing for the presence or amount of the IGS4 polypeptide expression in a sample derived from said subject.

26. (ORIGINAL) A method for identifying agonists to the IGS4 polypeptide of claim 17-24 comprising:

(a) contacting a cell which produces a IGS4 polypeptide with a test compound; and

(b) determining whether the test compound effects a signal generated by activation of the IGS4 polypeptide.

27. (ORIGINAL) An agonist identified by the method of claim 26.

28. (CURRENTLY AMENDED) A method for identifying agonists to the IGS4 neuromedin receptor protein, ~~preferably to the mammalian IGS4 neuromedin-receptor protein,~~ said protein exhibiting high affinity binding for neuromedin U, ~~preferably for neuromedin U-8, for neuromedin U-23 and/or for neuromedin U-25,~~ comprising:

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(a) contacting a cell which produces a IGS4 neuromedin receptor protein with a test compound; and

(b) determining whether the test compound effects a signal generated by activation of the IGS4 neuromedin receptor protein.

29. (CURRENTLY AMENDED) A method for identifying agonists to the IGS4 neuromedin receptor protein according to claim 28, wherein said agonists are effective with regard to at least one of disorders of the nervous system, including ~~the central nervous system (CNS) and the peripheral nervous system (PNS)~~, disorders of the gastrointestinal system, and/or disorders of the cardiovascular system, disorders of and/or of the skeletal muscle, and/or of disorders of the thyroid, ~~and/or also to~~ lung diseases, immunological diseases, and disorders of the genitourinary system.
30. (CURRENTLY AMENDED) An agonist identified by the method of claim 28 ~~or 29~~, preferably an agonist being effective with regard to ~~disorders of the nervous system, including the central nervous system (CNS) and the peripheral nervous system (PNS), disorders of the gastrointestinal system and/or of the cardiovascular system and/or of skeletal muscle and/or of the thyroid, and/or also to lung diseases, immunological diseases and disorders of the genitourinary system.~~
31. (CURRENTLY AMENDED) A method for identifying antagonists to the IGS4 polypeptide of claim 17-24 comprising:

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- (a) contacting a cell which produces a IGS4 polypeptide with an agonist;  
and  
(b) determining whether the signal generated by said agonist is diminished in the presence of a candidate compound.

32. (ORIGINAL) An antagonist identified by the method of claim 31.
33. (CURRENTLY AMENDED) A method for identifying antagonists to the IGS4 neuromedin receptor protein, ~~preferably to the mammalian IGS4 neuromedin receptor protein~~, said protein exhibiting high affinity binding for neuromedin U, ~~preferably for neuromedin U-8, for neuromedin U-23 and/or for neuromedin U-25~~, comprising:
- (a) contacting a cell which produces a IGS4 neuromedin receptor protein with an agonist; and  
(b) determining whether the signal generated by said agonist is diminished in the presence of a candidate compound.
34. (CURRENTLY AMENDED) A method for identifying antagonists to the IGS4 neuromedin receptor protein according to claim 33, wherein said antagonists are effective with regard at least one disorder chosen from disorders of ~~to disorders of~~ the nervous system, ~~including the central nervous system (CNS) and the peripheral nervous system (PNS)~~, disorders of the gastrointestinal system, ~~and/or of the cardiovascular system, and/or of the skeletal muscle, and/or of the thyroid, and/or also to the lung diseases, immune system, or immunological diseases and disorders of the genitourinary system.~~

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35. (CURRENTLY AMENDED) An antagonist identified by the method of claim 33 ~~or 34, preferably an antagonist being effective with regard to disorders of the nervous system, including the central nervous system (CNS) and the peripheral nervous system (PNS), disorders of the gastrointestinal system and/or of the cardiovascular system and/or of skeletal muscle and/or of the thyroid, and/or also to lung diseases, immunological diseases and disorders of the genitourinary system.~~
36. (CURRENTLY AMENDED) A recombinant host cell, or a membrane of a recombinant host cell produced by a the method of claim 16 ~~or a membrane thereof expressing wherein the host cell or membrane expresses an IGS4 polypeptide.~~
37. (ORIGINAL) A method of creating a genetically modified non-human animal comprising ~~the steps of:~~
- (a) ligating the coding portion of a nucleic acid molecule, consisting essentially of a nucleic acid sequence encoding a protein having the amino acid sequence SEQ ID NO: 2, SEQ ID NO: 4, SEQ NO: 6 or SEQ NO: 8 or the amino acid sequence encoded by the DNA insert contained in the deposit no. CBS102221 or the deposit no. CBS1 02222 at the Centraalbureau voor Schimmelcultures at Baarn the Netherlands or a biologically active portion of one of said sequences, to a regulatory sequence which is capable of driving high level gene expression or expression in a cell type in which the gene is not normally expressed in said animal; or

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(b) isolation and engineering the coding portion of a nucleic acid molecule, consisting essentially of a nucleic acid sequence encoding a protein having the amino acid sequence SEQ ID NO: 2, SEQ ID NO: 4, SEQ NO: 6 or SEQ NO: 8 or the amino acid sequence encoded by the DNA insert contained in the deposit no. CBS102221 or the deposit no. CBS102222 at the Centraalbureau voor Schimmelcultures at Baarn the Netherlands or a biologically active portion of one of said sequences, and reintroducing said sequence in the genome of said animal in such a way that the endogenous gene alleles, encoding a protein having the amino acid sequence SEQ ID NO: 2, SEQ ID NO: 4, SEQ NO: 6 or SEQ NO: 8 or the amino acid sequence encoded by the DNA insert contained in the deposit no. CBS102221 or the deposit no. CBS102222 at the Centraalbureau voor Schimmelcultures at Baarn the Netherlands or a biologically active portion of one of said sequences, are fully or partially inactivated.

38. (ORIGINAL) A method of determining whether a substance is a potential ligand of IGS4 receptor comprising:

(a) contacting cells expressing the receptor of ~~one of the claims 17-24~~ claim 17 or one of SEQ ID NO:2, SEQ ID NO:4 SEQ ID NO:6 and SEQ ID NO:8, or contacting a receptor membrane preparation comprising one of said receptors of ~~one of the claims 17-24~~ claim 17 or one of SEQ ID NO:2, SEQ ID NO:4 SEQ ID NO:6 and SEQ ID NO:8 with labeled neuromedin U in the presence and in the absence of the substance; and

(b) measuring the binding of neuromedin U to IGS4.

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39. (CURRENTLY AMENDED) A polypeptide according to ~~any of the claims 17-21,~~  
claim 17, wherein the further being characterized in that said polypeptide binds  
neuromedin U, ~~preferably neuromedin U-8, neuromedin U-23 and/or neuromedin~~  
U-25, and has showing at least an affinity of about at least  $\log EC_{50} = -6$ .
40. (CURRENTLY AMENDED) A polypeptide according to ~~any of the claims 17-21,~~  
claim 17, wherein the further being characterized in that said polypeptide binds  
neuromedin U, ~~preferably neuromedin U-8, neuromedin U-23 and/or neuromedin~~  
U-25, and has showing at least an affinity of at least about  $\log EC_{50} = -9$ .
41. (NEW) The isolated polynucleotide of claim 1, wherein the nucleotide sequence  
has at least 90% sequence identity over its entire length to the nucleotide  
sequence of (a) or (b) or (c).
42. (NEW) The isolated nucleotide sequence of claim 6, wherein the IGS4  
neuromedin receptor protein is a mammalian neuromedin receptor protein and  
neuromedin U is at least one of neuromedin U-8, neuromedin U-23, and  
neuromedin U-25.
43. (NEW) The isolated nucleotide sequence of claim 8, wherein the IGS4  
neuromedin receptor protein is a mammalian neuromedin receptor protein and  
neuromedin U is at least one of neuromedin U-8, neuromedin U-23, and  
neuromedin U-25.
44. (NEW) The isolated DNA or RNA molecule of claim 10, wherein the IGS4  
neuromedin receptor protein is a mammalian neuromedin receptor protein and

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neuromedin U is at least one of neuromedin U-8, neuromedin U-23, and neuromedin U-25.

45. (NEW) The isolated IGS4 polypeptide of claim 19, wherein the IGS4 neuromedin receptor protein is a mammalian neuromedin receptor protein and neuromedin U is at least one of neuromedin U-8, neuromedin U-23, and neuromedin U-25.
46. (NEW) The isolated IGS4 polypeptide of claim 21, wherein the IGS4 neuromedin receptor protein is a mammalian neuromedin receptor protein and neuromedin U is at least one of neuromedin U-8, neuromedin U-23, and neuromedin U-25.
47. (NEW) The method of treatment of claim 23, wherein the IGS4 neuromedin receptor protein is a mammalian neuromedin receptor protein and neuromedin U is at least one of neuromedin U-8, neuromedin U-23, and neuromedin U-25.
48. (NEW) The method of identifying agonists of claim 28, wherein the IGS4 neuromedin receptor protein is a mammalian neuromedin receptor protein and neuromedin U is at least one of neuromedin U-8, neuromedin U-23, and neuromedin U-25.
49. (NEW) The agonist of claim 30, wherein the agonist is effective with regard to at least one disorder chosen from disorders of the nervous system, ~~including the central nervous system (CNS) and peripheral nervous system (PNS)~~, the gastrointestinal system, the cardiovascular system, the skeletal muscle, the thyroid, the lung, the immune system, or the genitourinary system.

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50. (NEW) The method of identifying antagonists of claim 33, wherein the IGS4 neuromedin receptor protein is a mammalian neuromedin receptor protein and neuromedin U is at least one of neuromedin U-8, neuromedin U-23, and neuromedin U-25.
51. (NEW) The antagonist of claim 35, wherein the antagonist is effective with regard to disorders of at least one of the nervous system, ~~including the central nervous system (CNS) and peripheral nervous system (PNS)~~, the gastrointestinal system, the cardiovascular system, the skeletal muscle, the thyroid, the lung, the immune system, and the genitourinary system.
52. (NEW) The polypeptide of claim 39, wherein the neuromedin U is at least one of neuromedin U-8, neuromedin U-23, and neuromedin U-25.
53. (NEW) The polypeptide of claim 40, wherein the neuromedin U is at least one of neuromedin U-8, neuromedin U-23, and neuromedin U-25.
54. (NEW) A host cell comprising the expression system of claim 10.
55. (NEW) The method of identifying agonists to the IGS4 neuromedin receptor protein according to claim 29, wherein disorders of the nervous system are disorders of the central nervous system (CNS) or the peripheral nervous system (PNS).
56. (NEW) The method for identifying antagonists to the IGS4 neuromedin receptor protein according to claim 34, wherein disorders of the nervous system are

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disorders of the central nervous system (CNS) or the peripheral nervous system (PNS).

57. (NEW) The agonist of claim 49, wherein the disorders of the nervous system are disorders of the central nervous system (CNS) or peripheral nervous system (PNS).
58. (NEW) The antagonist of claim 51, wherein the disorders of the nervous system are disorders of the central nervous system (CNS) and peripheral nervous system (PNS).
59. (NEW) An agonist identified by the method of claim 29.
60. (NEW) An isolated IGS4 polypeptide comprising an amino acid sequence of a neuromedin receptor protein, said protein exhibiting high affinity binding for neuromedin U, said protein exhibiting expression in at least one of brain, skeletal muscle, cerebellum, testis, corpus callosum, spinal cord, substantia nigra, medulla, thalamus, caudate nucleus, pons, nucleus accumbens, fetal brain, stomach, heart, thyroid gland, lung, thymus, prostate, and trachea, and said amino acid sequence being selected from the group of amino acid sequences as defined in claim 18.
61. (NEW) An antibody immunospecific for the IGS4 polypeptide of claim 18.
62. (NEW) A method for the treatment of a subject in need of enhanced activity or expression of IGS4 polypeptide of claim 18 comprising at least one of:

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(a) administering to the subject a therapeutically effective amount of an agonist to said receptor;

(b) providing to the subject an isolated polynucleotide comprising a nucleotide sequence that has at least 80% identity to a nucleotide sequence encoding the IGS4 polypeptide of SEQ ID NO: 2, SEQ ID NO: 4, SEQ NO: 6 or SEQ NO: 8 or the polypeptide encoded by the DNA insert contained in the deposit no. CBS102221 or the deposit no. CBS102222 at the Centraalbureau voor Schimmelcultures at Baarn the Netherlands over its entire length; or a nucleotide sequence complementary to one of said nucleotide sequences in a form so as to effect production of said receptor activity in vivo; and,

(c) providing to the subject an isolated polynucleotide comprising a nucleotide sequence that encodes an IGS4 neuromedin receptor protein, said protein exhibiting high affinity binding for neuromedin U.

63. (NEW) A method for the treatment of a subject having need to inhibit activity or expression of IGS4 polypeptide of claim 18 comprising at least one of:

(a) administering to the subject a therapeutically effective amount of an antagonist to said receptor;

(b) administering to the subject a nucleic acid molecule that inhibits the expression of the nucleotide sequence encoding said receptor; and,

(c) administering to the subject a therapeutically effective amount of a polypeptide that competes with said receptor for its ligand.

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64. (NEW) A process for diagnosing a disease or a susceptibility to a disease in a subject related to expression or activity of the IGS4 polypeptide of claim 18 in a subject comprising at least one of:
- (a) determining the presence or absence of a mutation in the nucleotide sequence encoding said IGS4 polypeptide in the genome of said subject; and,
  - (b) analyzing for the presence or amount of the IGS4 polypeptide expression in a sample derived from said subject.
65. (NEW) A method for identifying agonists to the IGS4 polypeptide of claim 18 comprising:
- (a) contacting a cell which produces a IGS4 polypeptide with a test compound; and
  - (b) determining whether the test compound effects a signal generated by activation of the IGS4 polypeptide.
66. (NEW) A method for identifying antagonists to the IGS4 polypeptide of claim 18 comprising:
- (a) contacting a cell which produces a IGS4 polypeptide with an agonist; and
  - (b) determining whether the signal generated by said agonist is diminished in the presence of a candidate compound.
67. (NEW) A method of determining whether a substance is a potential ligand of IGS4 receptor comprising:

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(a) contacting cells expressing the receptor of claim 18 or one of SEQ ID NO:2, SEQ ID NO:4 SEQ ID NO:6 and SEQ ID NO:8, or contacting a receptor membrane preparation comprising one of said receptors of claim 18 or one of SEQ ID NO:2, SEQ ID NO:4 SEQ ID NO:6 and SEQ ID NO:8 with labeled neuromedin U in the presence and in the absence of the substance; and

(b) measuring the binding of neuromedin U to IGS4.

68. (NEW) A polypeptide according to, claim 18, wherein the polypeptide binds neuromedin U, and has an affinity of about at least  $\log EC_{50} = -6$ .
69. (NEW) A polypeptide according to, claim 18, wherein the polypeptide binds neuromedin U, and has an affinity of at least about  $\log EC_{50} = -9$ .

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